AMENDMENT

Subject matter to be added is in bold and underlined.

Subject matter to be deleted is in bold and strikethrough.

In the Claims:

Please enter rewritten Claims 1-12 and 19 and new Claims 28-31 as follows. Please cancel Claims 24-27 without prejudice or disclaimer.

This listing of claims will replace all prior versions and listings of claims in the application.

1. (Currently amended) A compound according to formula (I),

or a stereoisomer or a pharmaceutically-acceptable salt thereof, wherein:

X is -OH, -O(alkyl), -O(arylalkyl), $-NR_5(aryl)$, or $-NR_5(arylalkyl)$; wherein said aryl or arylalkyl are optionally substituted with one to two R_{25} ;

W is hydrogen or -(CR₇R₈)_a-H;

Z is a 5-membered heteroaryl group optionally substituted with 1-3 R₉, a five to six membered heterocyclo or cycloalkyl group optionally substituted with 1-3 R₉, a 9 to 10 membered bicyclic aryl or heteroaryl isoquinolyl optionally

substituted with 1-3 substituents selected from
$$R_9$$
 and/or R_{10} , or R_{10} , R_{10}

Z₁, Z₂ and Z₃ are independently N or CR₉;

 $R_1,\,R_2$ and R_3 are attached to any available carbon atom of phenyl ring A and are independently selected from hydrogen, halogen, cyano, nitro, $C_{1\text{-}10}$ alkyl, $C_{2\text{-}10}$ alkenyl, substituted $C_{1\text{-}10}$ alkyl, substituted $C_{2\text{-}10}$ alkenyl, $-C(=O)NR_{12}R_{13},$ $-OR_{12},$ $-CO_2R_{12},$ $-C(=O)R_{12},$ $-SR_{12},$ $-S(O)_tR_{15},$ $-NR_{12}R_{13},$ $-NR_{12}SO_2R_{15},$ $-NR_{14}SO_2NR_{12}R_{13},$ $-NR_{12}CO_2R_{13},$ $-NR_{12}C(=O)R_{13},$ $-NR_{14}C(=O)NR_{12}R_{13},$ $-SO_2NR_{12}R_{13},$ aryl, heteroaryl, cycloalkyl, and heterocyclo;

R₅ is hydrogen, C₁₋₄alkyl, NH₂, C₁₋₄alkylamino, hydroxy, or C₁₋₄alkoxy;

 R_7 and R_8 are independently selected from hydrogen, $-OR_{18}$, $-NR_{18}R_{19}$, $-NR_{18}SO_2R_{20}$, alkyl, alkenyl, substituted alkyl, substituted alkenyl, halogen, haloalkyl, haloalkoxy, cyano, nitro, alkylthio, -C(=O)H, acyl, $-CO_2H$, alkoxycarbonyl, sulfonamido, sulfonyl, and phenyl in turn optionally substituted with 1-3 of halogen, cyano, haloalkyl, haloalkoxy, nitro, hydroxy, C_{1-4} alkyl, C_{1-4} hydroxyalkyl, C_{1-4} alkoxy, amino, $NH(C_{1-4}$ alkyl), $N(C_{1-4}$ alkyl)₂, and/or C_{1-4} aminoalkyl;

 R_{97} and R_{10} and R_{11} are independently selected from hydrogen, halogen, alkyl, substituted alkyl, haloalkyl, haloalkoxy, cyano, nitro, $-S(O)_uR_{21}$, $-NR_{22}SO_2R_{21}$, $-C(=O)NR_{22}R_{23}$, $-OR_{22}$, $-CO_2R_{22}$, $-C(=O)R_{22}$, $-SR_{22}$, $-NR_{22}R_{23}$, $-NR_{22}CO_2R_{23}$, $-NR_{22}C(=O)R_{23}$, $-NR_{22}C(=O)NR_{23}R_{24}$, $-SO_2NR_{22}R_{23}$, $-NR_{22}SO_2NR_{23}R_{24}$, $-C(=NR_{22})NR_{23}R_{24}$, five or six membered heterocyclo or heteroaryl, phenyl, and C_{3-7} cycloalkyl, provided that R_{11} is not $-C(=NR_{22})NR_{23}R_{24}$ when W or W_1 is hydrogen; wherein when R_{97} or R_{10} or R_{11} is selected from heterocyclo, heteroaryl, phenyl, and C_{3-7} cycloalkyl, each of said cyclic groups in turn is optionally substituted with up to three of C_{1-4} alkyl, C_{1-4} alkoxy, C_{1-4} hydroxyalkyl, C_{1-4} aminoalkyl, halogen, hydroxy, haloalkyl, haloalkoxy, amino, C_{1-4} alkylamino, and/or cyano;

 R_{12} , R_{13} , R_{14} , R_{18} , R_{19} , R_{22} R_{23} , and R_{24} are independently selected from hydrogen, alkyl, substituted alkyl, alkenyl, substituted alkenyl, aryl, heteroaryl, cycloalkyl, and heterocyclo;

 R_{15} , R_{20} and R_{21} are independently selected from alkyl, substituted alkyl, alkenyl, substituted alkenyl, aryl, heteroaryl, cycloalkyl, and heterocyclo;

 R_{25} at each occurrence is selected from hydrogen, halogen, cyano, nitro, $C_{1\text{-}10}$ alkyl, $C_{2\text{-}10}$ alkenyl, substituted $C_{1\text{-}10}$ alkyl, substituted $C_{2\text{-}10}$ alkenyl, $-C(=O)NR_{12}R_{13}$, $-OR_{12}$, $-CO_2R_{12}$, $-C(=O)R_{12}$, $-SR_{12}$, $-S(O)_tR_{15}$, $-NR_{12}R_{13}$, $-NR_{12}SO_2R_{15}$, $-NR_{14}SO_2NR_{12}R_{13}$, $-NR_{12}CO_2R_{13}$, $-NR_{12}C(=O)R_{13}$, $-NR_{14}C(=O)NR_{12}R_{13}$, $-SO_2NR_{12}R_{13}$, aryl, heteroaryl, cycloalkyl, and heterocyclo; p is 1 or 2; q is 1, 2 or 3; t is 1 or 2; and u is 1 or 2;

provided that when Z is phenyl, pyridyl or pyridazinyl, R_9 , R_{10} and/or R_{11} are other than cyano or $-C(=NR_{22})NR_{23}R_{24}$.

2. (Currently amended) A compound according to claim 1, or a stereoisomer or a pharmaceutically-acceptable salt thereof, wherein the compound is of formula (Ia):

X is -OH, -O(phenyl) optionally substituted with one to two R_{25} , -O(benzyl) optionally substituted with one to two R_{25} , -NH(phenyl) optionally substituted with one to two R_{25} , or -NH(benzyl) optionally substituted with one to two R_{25} ;

W is hydrogen or $-(CH_2)_q$ -H;

Z is selected from a 5-membered heteroaryl group optionally substituted with 1-3 R₉, a five to six membered heterocyclo or cycloalkyl group optionally substituted

with 1-3 R₉, a 9 to 10 membered bicyclic aryl or heteroaryl isoquinolyl optionally

substituted with 1-3 substituents selected from R_9 and/or R_{10} , and R_{10} , R_{10}

 Z_1 , Z_2 and Z_3 are independently N or CR₉ and at least one of Z_1 , Z_2 and Z_3 is N;

 $R_1 \text{ and } R_2 \text{ are independently selected from hydrogen, halogen, cyano, nitro,} \\ C_{1\text{-}10}\text{alkyl}, C_{2\text{-}10}\text{alkenyl, substituted } C_{1\text{-}10}\text{alkyl, substituted } C_{2\text{-}10}\text{alkenyl,} \\ -C(=O)\text{NR}_{12}\text{R}_{13}, -O\text{R}_{12}, -CO_2\text{R}_{12}, -C(=O)\text{R}_{12}, -S\text{R}_{12}, -S(O)_{t}\text{R}_{15}, -N\text{R}_{12}\text{R}_{13}, \\ -N\text{R}_{12}\text{SO}_2\text{R}_{15}, -N\text{R}_{14}\text{SO}_2\text{NR}_{12}\text{R}_{13}, -N\text{R}_{12}\text{CO}_2\text{R}_{13}, -N\text{R}_{12}\text{C}(=O)\text{R}_{13}, \\ -N\text{R}_{14}\text{C}(=O)\text{NR}_{12}\text{R}_{13}, -S\text{O}_2\text{NR}_{12}\text{R}_{13}, \text{aryl, heteroaryl, cycloalkyl, and heterocyclo;} \\ \\$

 R_{97} and R_{10} and R_{11} are independently selected from hydrogen, halogen, alkyl, substituted alkyl, haloalkyl, haloalkoxy, cyano, nitro, $-S(O)_uR_{21}$, $-NR_{22}SO_2R_{21}$, $-C(=O)NR_{22}R_{23}$, $-OR_{22}$, $-CO_2R_{22}$, $-C(=O)R_{22}$, $-SR_{22}$, $-NR_{22}R_{23}$, $-NR_{22}CO_2R_{23}$, $-NR_{22}C(=O)R_{23}$, $-NR_{22}C(=O)NR_{23}R_{24}$, $-SO_2NR_{22}R_{23}$, $-NR_{22}SO_2NR_{23}R_{24}$, $-C(=NR_{22})NR_{23}R_{24}$, five or six membered heterocyclo or heteroaryl, phenyl, and C_3 . $-C(=NR_{22})NR_{23}R_{24}$, when $-C(=NR_{22})NR_{23}R_{24}$ when $-C(=NR_{22})NR_{23}$

 R_{12} , R_{13} , R_{14} , R_{18} , R_{19} , R_{22} R_{23} , and R_{24} are independently selected from hydrogen, alkyl, substituted alkyl, alkenyl, substituted alkenyl, aryl, heteroaryl, cycloalkyl, and heterocyclo;

 R_{15} , R_{20} and R_{21} are independently selected from alkyl, substituted alkyl, alkenyl, substituted alkenyl, aryl, heteroaryl, cycloalkyl, and heterocyclo; R_{16} is alkyl, substituted alkyl, alkenyl, substituted alkenyl, aryl, heteroaryl, cycloalkyl, or heterocyclo;

p is 1 or 2;

q is 1, 2 or 3; and

u is 1 or 2;

provided that when Z is phenyl, pyridyl or pyridazinyl, R_9 , R_{10} and/or R_{11} are other than cyano or $-C(=NR_{22})NR_{22}R_{24}$.

3. (Currently amended) A compound according to claim 2, wherein:

X is selected from -OH, -O(phenyl), -O(benzyl), -NH(phenyl), and wherein each phenyl or benzyl group is optionally substituted with one to two R_{25} ,

W is hydrogen or $-(CH_2)_a$ -H;

Z is selected from the group:

$$(R_9)_s$$

$$(R_9)_s$$
and
$$HN$$

R₁ and R₂ are OR₁₂;

 R_9 is selected from hydrogen, halogen, alkyl, substituted alkyl, haloalkyl, haloalkoxy, cyano, nitro, -S(O)_uR₂₁, -NR₂₂SO₂R₂₁, -C(=O)NR₂₂R₂₃, -OR₂₂, -CO₂R₂₂, -C(=O)R₂₂, -SR₂₂, -NR₂₂R₂₃, -NR₂₂CO₂R₂₃, -NR₂₂C(=O)R₂₃, -NR₂₂C(=O)NR₂₃R₂₄, -SO₂NR₂₂R₂₃, -NR₂₂SO₂NR₂₃R₂₄, five or six membered heterocyclo or heteroaryl, phenyl, and C₃₋₇cycloalkyl;

R₁₂, R₂₃ and R₂₄ are selected from hydrogen, alkyl, substituted alkyl, alkenyl, substituted alkenyl, aryl, heteroaryl, cycloalkyl, or heterocyclo;

R₂₁ is selected from alkyl, substituted alkyl, alkenyl, substituted alkenyl, aryl, heteroaryl, cycloalkyl, and heterocyclo;

 R_{25} at each occurrence is selected from $C_{1\text{-}4}$ alkyl, $C_{1\text{-}4}$ alkoxy, $C_{1\text{-}4}$ hydroxyalkyl, $C_{1\text{-}4}$ aminoalkyl, halogen, hydroxy, haloalkyl, haloalkoxy, amino, $C_{1\text{-}4}$ alkylamino, and/or cyano;

q is 1, 2 or 3;

s is 0, 1, or 2; and

u is 1 or 2;

provided that when Z is phenyl, R_9 and/or R_{11} are other than eyano or $-C(=NR_{22})NR_{22}R_{24}$.

4. (Currently amended) A compound according to claim 1, or a stereoisomer or a pharmaceutically-acceptable salt thereof, wherein the compound is of formula (Ib),

$$OR_{12b}$$
 OR_{12a}
 OR_{12a}

wherein:

X is selected from -O(phenyl), -O(benzyl), and -NH(phenyl) -NH(benzyl), wherein each group X is optionally substituted with one to two R_{25} ,

W is hydrogen or $-(CH_2)_q$ -H;

Z is selected from the group:

$$(R_9)_s \qquad (R_9)_s \qquad \text{and} \qquad (R_9)_s \qquad \vdots$$

 R_9 is independently selected from hydrogen, halogen, alkyl, aminoalkyl, hydroxyalkyl, haloalkyl, haloalkoxy, alkoxy, cyano, nitro, alkylamino, alkylthio, thioalkyl, $-C(=O)NH_2$, $-C(=O)NH(C_{1-4}alkyl)$, $-C(=O)N(C_{1-4}alkyl)_2$, five or six membered heterocyclo or heteroaryl, phenyl, and C_{3-7} cycloalkyl;

R_{12a} and R_{12b} are independently selected from hydrogen, alkyl, substituted alkyl, phenyl, and benzyl;

 R_{25} at each occurrence is selected from $C_{1\text{-}4}$ alkyl, $C_{1\text{-}4}$ alkoxy,

 $C_{1\text{-}4}$ hydroxyalkyl, $C_{1\text{-}4}$ aminoalkyl, halogen, hydroxy, haloalkyl, haloalkoxy, amino, $C_{1\text{-}4}$ alkylamino, and/or cyano;

p is 1 or 2; and

s is 0, 1 or 2;

provided that when Z is phenyl, R_9 and/or R_{11} -are other than cyano or $-C(=NR_{22})NR_{23}R_{24}.$

5. (Currently amended) A compound according to claim 1, or a stereoisomer or a pharmaceutically-acceptable salt thereof, wherein Z is selected from:

Z₅ is fused to ring A comprising the common carbon atom C* and is selected from:

 Z_6 is fused to ring A comprising the common carbon atom C* and is

Z₇ is fused to ring A comprising the common carbon atom C* and is selected from:

 Z_8 is fused to ring B-comprising the common nitrogen atom N* and is selected from

$$(R_9)_r \qquad (R_9)_r \qquad (R_9)_r \qquad N \qquad (R_9)_r \qquad N \qquad (R_9)_r \qquad (R_9)_$$

Zo is CH or N;

r is 0, 1, or 2; and

s is 0, 1, 2, or 3.

6. (Currently amended) A compound according to claim 1, or a stereoisomer or a pharmaceutically-acceptable salt, hydrate or prodrug thereof, wherein Z is selected from:

- 7. (Currently amended) A compound according to claim 1, or a stereoisomer or a pharmaceutically-acceptable salt, hydrate or prodrug thereof, wherein R_1 and R_2 are OR_{12} .
- 8. (Currently amended) A compound according to claim 7, or a stereoisomer or a pharmaceutically acceptable salt, hydrate or prodrug thereof, wherein R_{12} is C_{1-6} alkyl, phenyl, or benzyl optionally substituted with one to two of halogen, cyano, haloalkyl, haloalkoxy, nitro, hydroxy, C_{1-4} alkyl, C_{1-4} hydroxyalkyl, C_{1-4} alkoxy, amino, NH(C_{1-4} alkyl), and N(C_{1-4} alkyl)₂.
- 9. (Currently amended) A compound according to claim 8, or a stereoisomer or a pharmaceutically-acceptable salt, hydrate or prodrug thereof, wherein W is hydrogen.
- 10. (Currently amended) A compound according to claim 9, or a stereoisomer or a pharmaceutically-acceptable salt, hydrate or prodrug thereof, wherein X is NH(phenyl), or NH(benzyl), SO₂alkyl, or SO₂(phenyl) optionally substituted with one to two of C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄hydroxyalkyl, C₁₋₄aminoalkyl, halogen, hydroxy, haloalkyl, haloalkoxy, amino, C₁₋₄alkylamino, and/or cyano.
- 11. (Currently amended) A compound having the formula (Ib),

or a stereoisomer or a pharmaceutically-acceptable salt thereof, wherein:

X is selected from -O(phenyl) optionally substituted with one to two R_{25} , -O(benzyl) optionally substituted with one to two R_{25} , -NH(phenyl) optionally substituted with one to two R_{25} , and -NH(phenylalkyl) optionally substituted with one to two R_{25} ;

W is hydrogen or $-(CH_2)_q$ -H;

Z is selected from:

Z₁, Z₂ and Z₃ are selected from N and CR₉;

Z₄ is fused to ring A comprising the common carbon atom C* and is

 Z_5 is fused to ring A comprising the common carbon atom C* and is selected from:

Z₆ is fused to ring A comprising the common carbon atom C* and is

 Z_7 is fused to ring A comprising the common carbon atom C^* and is selected from:

 Z_{8} is fused to ring B comprising the common nitrogen atom N* and is selected from

Zo is CH or N;

 R_9 **and R_{10} are** <u>is</u> independently selected from hydrogen, halogen, alkyl, substituted alkyl, haloalkyl, haloalkoxy, cyano, nitro, $-S(O)_uR_{21}$, $-NR_{22}SO_2R_{21}$, $-C(=O)NR_{22}R_{23}$, $-OR_{22}$, $-CO_2R_{22}$, $-C(=O)R_{22}$, $-SR_{22}$, $-NR_{22}R_{23}$, $-NR_{22}CO_2R_{23}$, $-NR_{22}C(=O)R_{23}$, $-NR_{22}C(=O)NR_{23}R_{24}$, $-SO_2NR_{22}R_{23}$, $-NR_{22}SO_2NR_{23}R_{24}$, $-C(=NR_{22})NR_{23}R_{24}$, five or six membered heterocyclo or heteroaryl, phenyl, and C_{3-7} cycloalkyl, provided that R_9 **and R_{10} are** <u>is</u> not $-C(=NR_{22})NR_{23}R_{24}$ when W is hydrogen; wherein when R_9 **or** R_{10} is independently selected from heterocyclo, heteroaryl, phenyl, and C_{3-7} cycloalkyl, each of said cyclic groups in turn is optionally substituted with up to three of C_{1-4} alkyl, C_{1-4} alkoxy, C_{1-4} hydroxyalkyl, C_{1-4} aminoalkyl, halogen, hydroxy, haloalkyl, haloalkoxy, amino, C_{1-4} alkylamino, and/or cyano;

 R_{12} , R_{12a} , R_{12b} , R_{22} R_{23} , and R_{24} are independently selected from hydrogen, alkyl, substituted alkyl, alkenyl, substituted alkenyl, aryl, heteroaryl, cycloalkyl, and heterocyclo;

R₂₁ is selected from alkyl, substituted alkyl, alkenyl, substituted alkenyl, aryl, heteroaryl, cycloalkyl, and heterocyclo;

 R_{25} at each occurrence is selected from $C_{1\text{-}4}$ alkyl, $C_{1\text{-}4}$ alkoxy, $C_{1\text{-}4}$ hydroxyalkyl, $C_{1\text{-}4}$ aminoalkyl, halogen, hydroxy, haloalkyl, haloalkoxy, amino, $C_{1\text{-}4}$ alkylamino, and/or cyano;

p is 1 or 2;
q is 1, 2 or 3;
r is 0, 1, or 2;
s is 0, 1, 2, or 3;
t is 1 or 2; and
u is 1 or 2.

12. (Currently amended) A compound according to claim 11, or a stereoisomer or a pharmaceutically-acceptable salt thereof, wherein Z is selected from

13. (Original) A compound according to claim 1, wherein:

X is NR₅(benzyl) optionally substituted with one to two R₂₅;

W is hydrogen;

$$Z$$
 is $(R_9)_s$; and

 R_{25} at each occurrence is selected from halogen, cyano, nitro, C_{1-10} alkyl,

 C_{2-10} alkenyl, substituted C_{1-10} alkyl, substituted C_{2-10} alkenyl, $-C(=O)NR_{12}R_{13}$, $-OR_{12}$,

$$-\mathrm{CO_2R_{12}}, -\mathrm{C(=O)R_{12}}, -\mathrm{SR_{12}}, -\mathrm{S(O)_tR_{15}}, -\mathrm{NR_{12}R_{13}}, -\mathrm{NR_{12}SO_2R_{15}},$$

$$-NR_{14}SO_2NR_{12}R_{13},-NR_{12}CO_2R_{13},-NR_{12}C(=O)R_{13},-NR_{14}C(=O)NR_{12}R_{13},\\$$

-SO₂NR₁₂R₁₃, aryl, heteroaryl, cycloalkyl, and heterocyclo.

14. (Original) A compound according to claim 13, wherein:

15. (Original) A compound according to claim 13, wherein:

16. (Original) A compound according to claim 1, wherein:

X is OH;

W is hydrogen; and

17. (Original) A compound according to claim 16, wherein:

18. (Original) A compound according to claim 16, wherein:

- 19. (Currently amended) A compound according to claim 1, wherein the compound is selected from the group:
- 2 (4 Aminomethyl-phenylamino) N-benzyl-2 (3-ethoxy-4-isopropoxy-phenyl) acetamide;
- 7-{[Carboxy-(3-ethoxy-4-isopropoxy-phenyl)-methyl]-amino}-3,4-dihydro-1H-isoquinoline 2-earboxylicacid tert-butyl ester;
- [3-(tert-Butoxycarbonylamino-methyl)-phenylamino]-(3-ethoxy-4-isopropoxy-phenyl)-acetic acid;

(1-Amino-isoquinolin-6-ylamino)-(3-ethoxy-4-isopropoxy-phenyl)-acetic acid; and

- 2-(1-Amino-isoquinolin-6-ylamino)-N-benzyl-2-(3-ethoxy-4-isopropoxy-phenyl)-acetamide; or a stereoisomer or a pharmaceutically-acceptable salt thereof.
- 20. (Original) A pharmaceutical composition, comprising: a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 1, or a stereoisomer or a pharmaceutically-acceptable salt thereof.
- 21. (Original) A method for treating a thromboembolic disorder, comprising: administering to a patient in need thereof a therapeutically effective amount of a compound of Claim 1, or a stereoisomer or a pharmaceutically acceptable salt thereof.
- 22. (Original) A method according to Claim 21, wherein the thromboembolic disorder is selected from the group consisting of arterial cardiovascular thromboembolic disorders, venous cardiovascular thromboembolic disorders, and thromboembolic disorders in the chambers of the heart.
- 23. (Original) A method according to Claim 21, wherein the thromboembolic disorder is selected from unstable angina, an acute coronary syndrome, first myocardial infarction, recurrent myocardial infarction, ischemic sudden death, transient ischemic attack, stroke, atherosclerosis, peripheral occlusive arterial disease, venous thrombosis, deep vein thrombosis, thrombophlebitis, arterial embolism, coronary arterial thrombosis, cerebral arterial thrombosis, cerebral embolism, kidney embolism, pulmonary embolism, and thrombosis resulting from (a) prosthetic valves or other implants, (b) indwelling catheters, (c) stents, (d) cardiopulmonary bypass, (e) hemodialysis, or (f) other procedures in which blood is exposed to an artificial surface that promotes thrombosis.

24-27. (Canceled)

28. (New) A pharmaceutical composition, comprising: a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 2, or a stereoisomer or a pharmaceutically-acceptable salt thereof.

29. (New) A pharmaceutical composition, comprising: a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 3, or a stereoisomer or a pharmaceutically-acceptable salt thereof.

30. (New) A pharmaceutical composition, comprising: a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 4, or a stereoisomer or a pharmaceutically-acceptable salt thereof.

31. (New) A method for treating thrombosis, comprising: administering to a patient in need thereof a therapeutically effective amount of a compound of Claim 1, or a stereoisomer or a pharmaceutically acceptable salt thereof.